

Supplementary Data

Title

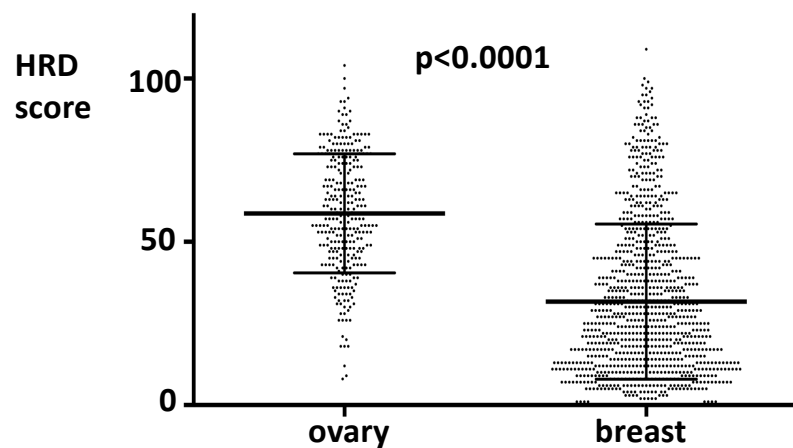
Homologous recombination deficiency status-based classification of high-grade serous ovarian carcinoma

Authors

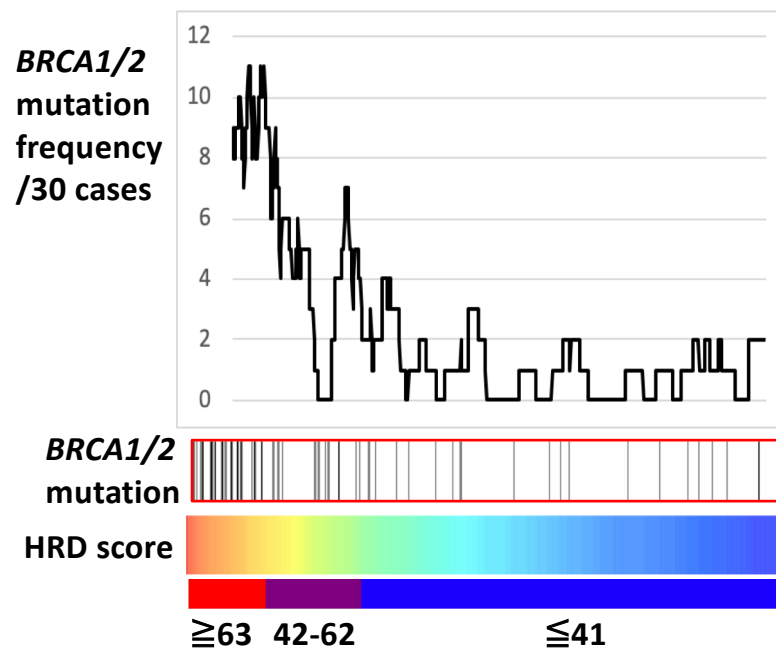
Hisamitsu Takaya, Hidekatsu Nakai, Shiro Takamatsu, Masaki Mandai, Noriomi Matsumura

Supplementary Figure 1

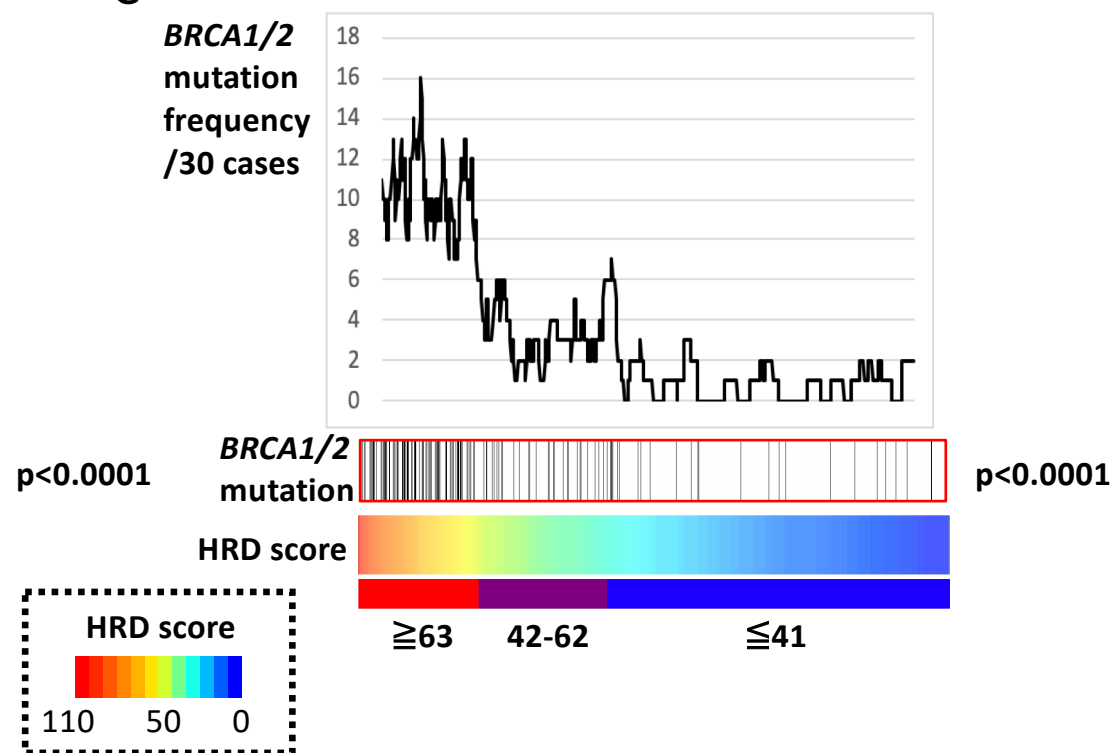
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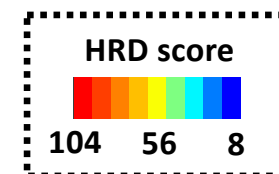
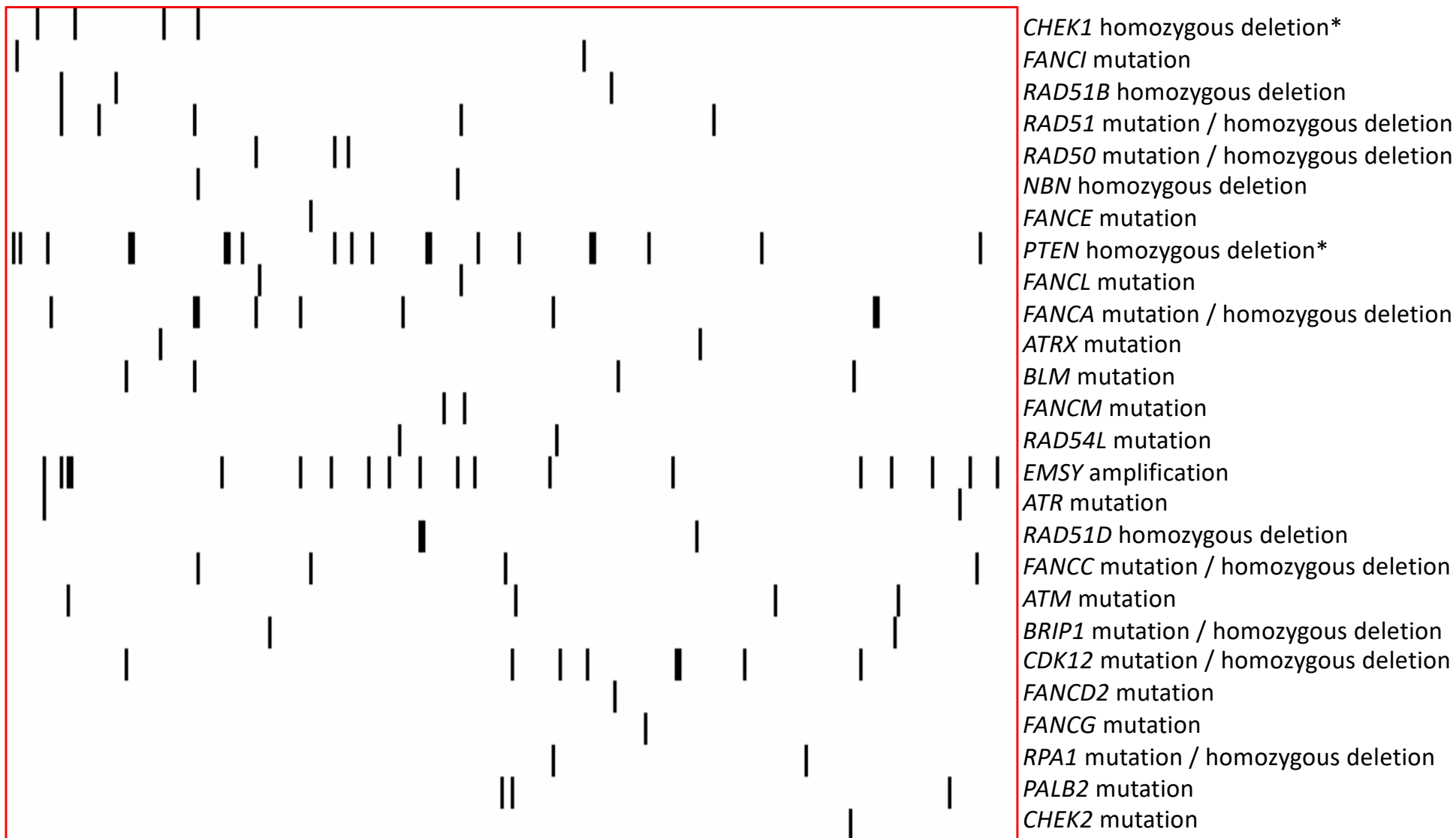
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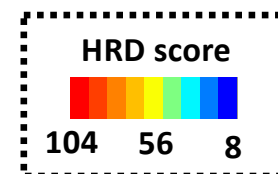
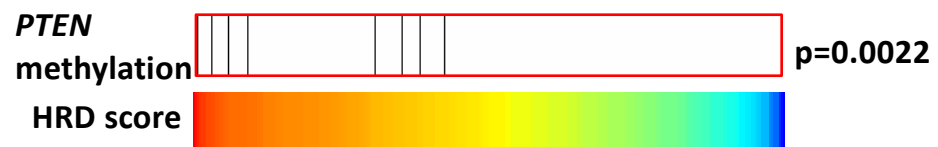
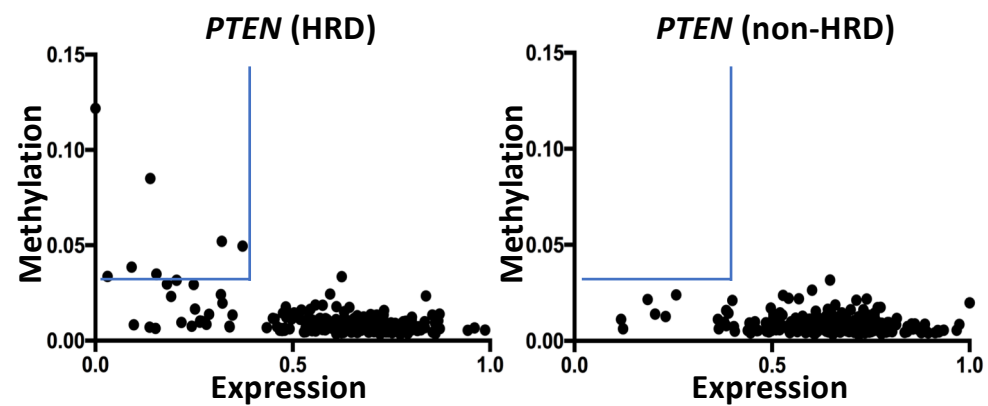
C



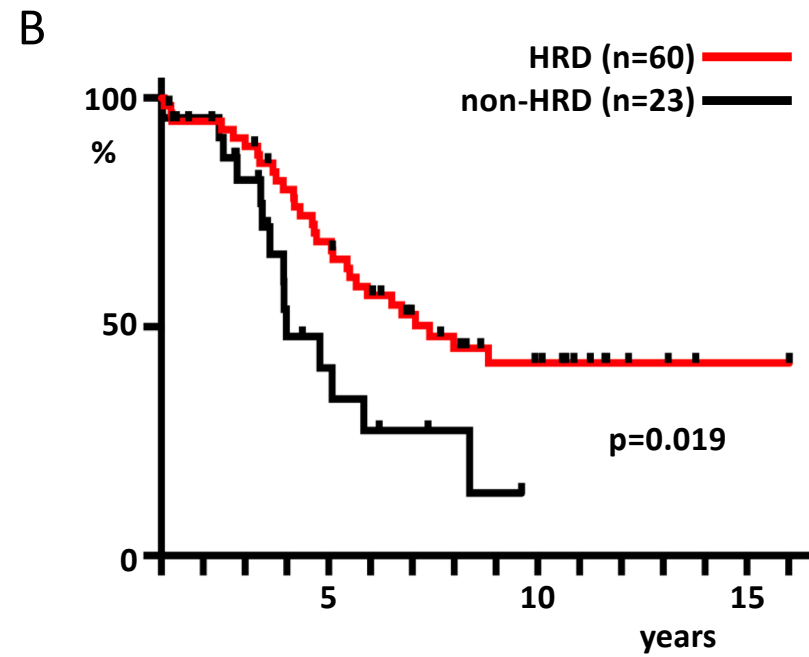
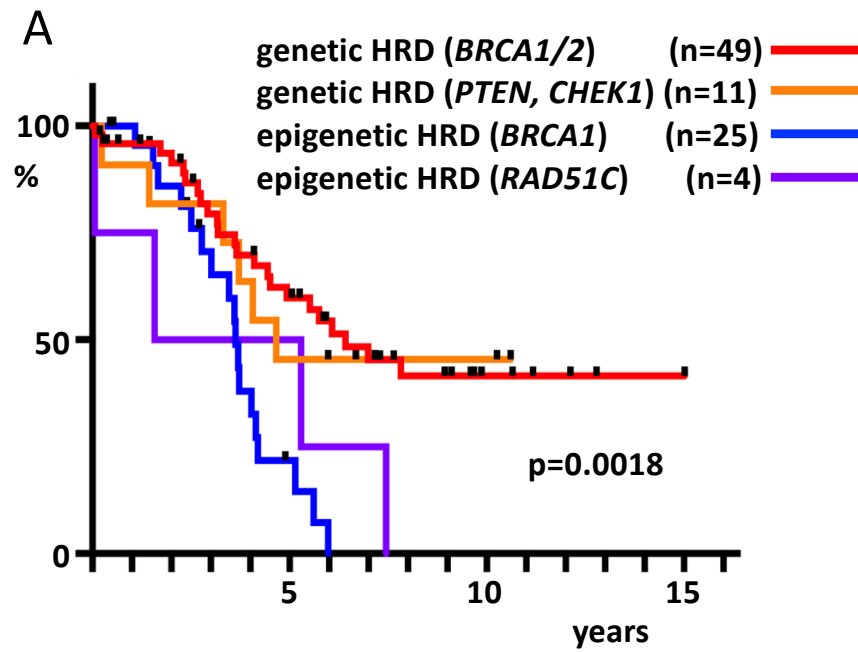
Supplementary Figure 2



Supplementary Figure 3



Supplementary Figure 4



Supplementary Figure Legends

Supplementary Figure 1. Analysis of the breast cancer dataset. A) Comparison of HRD scores. TCGA dataset samples with both exome sequencing and SNP array data (ovary: n = 296; breast: n = 955). B) Distribution of breast cancer *BRCA1/2* mutation cases. The upper graph shows the frequency of *BRCA* mutation cases; TCGA data were sorted by HRD scores using window sliding for 30 cases. The black bars in the central box represent the *BRCA* mutation cases; the higher the HRD score, the more enriched the *BRCA* mutations. Sections of HRD scores ≥ 63 , $42 - 62$, and $41 \geq$ are shown by colored bars. B) In all, 1251 cases (breast cancer: 955 cases, ovarian cancer: 296 cases) were sorted by HRD score to show the distribution of *BRCA* mutation cases. *BRCA1/2* mutation cases have a bimodal distribution, with an apparent cutoff ≥ 42 as well as ≥ 63 .

Supplementary Figure 2. Relationship of HRD scores and HRR pathway gene alterations other than *BRCA1/2*. Gene alterations are sorted by average HRD scores. Only *CHEK1* homozygous deletions and *PTEN* homozygous deletions were significantly enriched in the cases with high HRD scores (*; $p=0.0038$ and 0.035 , respectively).

Supplementary Figure 3. Analysis of *PTEN* methylation. X-axis: average values of three *PTEN* mRNA expression probes normalized to 0 – 1. Y-axis: β value of cg21573601. *PTEN* methylation cases are surrounded by a blue line (upper-left region). The frequency of *PTEN* methylation differs substantially between HRD cases (left) and non-HRD cases (right); β values are generally low.

Supplementary Figure 4. A) Comparison of HRD cases in OS. Genetic HRD cases were divided into those with *BRCA1/2* mutations and those with *CHEK1* or *PTEN* homozygous deletions; epigenetic HRD cases were divided into *BRCA1* methylation cases and *RAD51C* methylation cases. B) Cases with genetic factors (*BRCA1/2* methylations, *CHEK1* homozygous deletions, and *PTEN* homozygous deletions) were divided into HRD and non-HRD groups for comparison of OS.

[illegible]

Appendix 1: List of 1000 random numbers

Index	Value	Index	Value	Index	Value	Index	Value	Index	Value
1	0.1234	501	0.8765	991	0.5432	1	0.1234	501	0.8765
2	0.2345	502	0.7654	992	0.6543	2	0.2345	502	0.7654
3	0.3456	503	0.6543	993	0.7654	3	0.3456	503	0.6543
4	0.4567	504	0.5432	994	0.8765	4	0.4567	504	0.5432
5	0.5678	505	0.4321	995	0.9876	5	0.5678	505	0.4321
6	0.6789	506	0.3210	996	0.0987	6	0.6789	506	0.3210
7	0.7890	507	0.2109	997	0.1098	7	0.7890	507	0.2109
8	0.8901	508	0.1098	998	0.2109	8	0.8901	508	0.1098
9	0.9012	509	0.0987	999	0.3210	9	0.9012	509	0.0987
10	0.0123	510	0.9876	1000	0.4321	10	0.0123	510	0.9876
11	0.1123	511	0.8876			11	0.1123	511	0.8876
12	0.2123	512	0.7876			12	0.2123	512	0.7876
13	0.3123	513	0.6876			13	0.3123	513	0.6876
14	0.4123	514	0.5876			14	0.4123	514	0.5876
15	0.5123	515	0.4876			15	0.5123	515	0.4876
16	0.6123	516	0.3876			16	0.6123	516	0.3876
17	0.7123	517	0.2876			17	0.7123	517	0.2876
18	0.8123	518	0.1876			18	0.8123	518	0.1876
19	0.9123	519	0.0876			19	0.9123	519	0.0876
20	0.0234	520	0.9765			20	0.0234	520	0.9765
21	0.1234	521	0.8765			21	0.1234	521	0.8765
22	0.2234	522	0.7765			22	0.2234	522	0.7765
23	0.3234	523	0.6765			23	0.3234	523	0.6765
24	0.4234	524	0.5765			24	0.4234	524	0.5765
25	0.5234	525	0.4765			25	0.5234	525	0.4765
26	0.6234	526	0.3765			26	0.6234	526	0.3765
27	0.7234	527	0.2765			27	0.7234	527	0.2765
28	0.8234	528	0.1765			28	0.8234	528	0.1765
29	0.9234	529	0.0765			29	0.9234	529	0.0765
30	0.0345	530	0.9654			30	0.0345	530	0.9654
31	0.1345	531	0.8654			31	0.1345	531	0.8654
32	0.2345	532	0.7654			32	0.2345	532	0.7654
33	0.3345	533	0.6654			33	0.3345	533	0.6654
34	0.4345	534	0.5654			34	0.4345	534	0.5654
35	0.5345	535	0.4654			35	0.5345	535	0.4654
36	0.6345	536	0.3654			36	0.6345	536	0.3654
37	0.7345	537	0.2654			37	0.7345	537	0.2654
38	0.8345	538	0.1654			38	0.8345	538	0.1654
39	0.9345	539	0.0654			39	0.9345	539	0.0654
40	0.0456	540	0.9543			40	0.0456	540	0.9543
41	0.1456	541	0.8543			41	0.1456	541	0.8543
42	0.2456	542	0.7543			42	0.2456	542	0.7543
43	0.3456	543	0.6543			43	0.3456	543	0.6543
44	0.4456	544	0.5543			44	0.4456	544	0.5543
45	0.5456	545	0.4543			45	0.5456	545	0.4543
46	0.6456	546	0.3543			46	0.6456	546	0.3543
47	0.7456	547	0.2543			47	0.7456	547	0.2543
48	0.8456	548	0.1543			48	0.8456	548	0.1543
49	0.9456	549	0.0543			49	0.9456	549	0.0543
50	0.0567	550	0.9432			50	0.0567	550	0.9432
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53	0.3567	553	0.6432			53	0.3567	553	0.6432
54	0.4567	554	0.5432			54	0.4567	554	0.5432
55	0.5567	555	0.4432			55	0.5567	555	0.4432
56	0.6567	556	0.3432			56	0.6567	556	0.3432
57	0.7567	557	0.2432			57	0.7567	557	0.2432
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63	0.3678	563	0.6321			63	0.3678	563	0.6321
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138	0.9345	638	0.0654			138	0.9345	638	0.0654
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2000	1001	1002	1003	1004	1005	1006	1007	1008	1009	1010	1011	1012	1013	1014	1015	1016	1017	1018	1019	1020	1021	1022	1023	1024	1025	1026	1027	1028	1029	1030	1031	1032	1033	1034	1035	1036	1037	1038	1039	1040	1041	1042	1043	1044	1045	1046	1047	1048	1049	1050	1051	1052	1053	1054	1055	1056	1057	1058	1059	1060	1061	1062	1063	1064	1065	1066	1067	1068	1069	1070	1071	1072	1073	1074	1075	1076	1077	1078	1079	1080	1081	1082	1083	1084	1085	1086	1087	1088	1089	1090	1091	1092	1093	1094	1095	1096	1097	1098	1099	1100											
2001	1101	1102	1103	1104	1105	1106	1107	1108	1109	1110	1111	1112	1113	1114	1115	1116	1117	1118	1119	1120	1121	1122	1123	1124	1125	1126	1127	1128	1129	1130	1131	1132	1133	1134	1135	1136	1137	1138	1139	1140	1141	1142	1143	1144	1145	1146	1147	1148	1149	1150	1151	1152	1153	1154	1155	1156	1157	1158	1159	1160	1161	1162	1163	1164	1165	1166	1167	1168	1169	1170	1171	1172	1173	1174	1175	1176	1177	1178	1179	1180	1181	1182	1183	1184	1185	1186	1187	1188	1189	1190	1191	1192	1193	1194	1195	1196	1197	1198	1199	1200											
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